Test Type: CHRONIC

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH **AVERAGE SEVERITY GRADES[b]**

1,2-Dibromo-2,4-dicyanobutane

**CAS Number:** 35691-65-7

F\_1.Rev.1\_M3

C Number: C97003D

Lock Date: 01/31/2005

**Cage Range:** ALL

**Date Range:** ALL

**Reasons For Removal:** ALL

**Removal Date Range:** ALL

**Treatment Groups:** Include ALL

**TDMSE Version:** 1.8.0 Date Report Regsted: 08/21/2007

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Date Report Reqsted: 08/21/2007

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

| B6C3F1 MICE MALE                        | 0 MG/KG            | 0.6 MG/KG | 2 MG/KG  | 6 MG/KG  |  |
|---|--------------------|-----------|----------|----------|--|
| Disposition Summary                     |                    |           |          |          |  |
| Animals Initially in Study              | 50                 | 50        | 50       | 50       |  |
| Early Deaths                            | 00                 | 00        | 00       | 00       |  |
| Accidently Killed                       |                    | 1         |          |          |  |
| Moribund Sacrifice                      | 8                  | 12        | 8        | 7        |  |
| Natural Death                           | 7                  | 7         | 3        | 3        |  |
| Survivors                               |                    |           |          |          |  |
| Moribund Sacrifice                      | 1                  | 00        | 00       | 40       |  |
| Terminal Sacrifice                      | 34<br>50           | 30<br>50  | 39<br>50 | 40<br>50 |  |
| Animals Examined Microscopically        | 50                 | 50        | 50       | 50       |  |
| ALIMENTARY SYSTEM                       |                    |           |          |          |  |
| Esophagus                               | (50)               | (50)      | (50)     | (50)     |  |
| Gallbladder                             | (49)               | (49)      | (48)     | (49)     |  |
| Inflammation                            | 3 [1.0]            | ( - /     | 3 [1.0]  | 2 [1.0]  |  |
| Intestine Large, Cecum                  | (50)               | (50)      | (50)     | (50)     |  |
| Hyperplasia, Lymphoid                   |                    |           |          | 1 [3.0]  |  |
| Intestine Large, Colon                  | (50)               | (50)      | (50)     | (50)     |  |
| Intestine Large, Rectum                 | (50)               | (50)      | (50)     | (50)     |  |
| Intestine Small, Duodenum               | (50)               | (50)      | (50)     | (50)     |  |
| Ectopic Tissue                          | (=0)               | (40)      | 1 [3.0]  | (50)     |  |
| Intestine Small, Ileum                  | (50)               | (49)      | (50)     | (50)     |  |
| Inflammation<br>Ulcer                   | 1 [3.0]            |           |          |          |  |
| Peyer's Patch, Hyperplasia, Lymphoid    | 1 [3.0]<br>1 [4.0] |           |          |          |  |
| Intestine Small, Jejunum                | (50)               | (50)      | (50)     | (50)     |  |
| Peyer's Patch, Hyperplasia, Lymphoid    | (50)               | 1 [4.0]   | 4 [3.3]  | 2 [3.0]  |  |
| Liver                                   | (50)               | (50)      | (50)     | (50)     |  |
| Basophilic Focus                        | 1                  | 4         | 5        | 4        |  |
| Clear Cell Focus                        | 34                 | 22        | 29       | 24       |  |
| Eosinophilic Focus                      | 21                 | 9         | 16       | 16       |  |
| Fatty Change, Focal                     | 11 [1.4]           |           | 4 [1.5]  | 6 [1.7]  |  |
| Fatty Change, Diffuse                   | 19 [1.2]           | 15 [1.1]  | 11 [1.0] | 14 [1.0] |  |
| Hematopoietic Cell Proliferation        | 3 [1.3]            | 9 [1.3]   | 2 [1.0]  | 1 [1.0]  |  |
| Infarct                                 | 1 [4.0]            |           |          |          |  |
| Infiltration Cellular, Mononuclear Cell | 1 [2.0]            |           |          |          |  |

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Test Type: CHRONIC

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Date Report Reqsted: 08/21/2007

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

Lab: BAT

| 6C3F1 MICE MALE                         | 0 MG/KG  | 0.6 MG/KG | 2 MG/KG  | 6 MG/KG  |  |
|---|----------|-----------|----------|----------|--|
| Inflammation                            | 31 [1.0] | 19 [1.1]  | 27 [1.0] | 28 [1.0] |  |
| Mineralization                          | 1 [2.0]  |           |          |          |  |
| Mixed Cell Focus                        | 14       | 9         | 13       | 12       |  |
| Necrosis                                | 3 [1.7]  | 9 [1.9]   | 5 [1.8]  | 7 [2.0]  |  |
| Pigmentation                            | 4 [1.8]  | 3 [1.3]   |          |          |  |
| Tension Lipidosis                       | 3        | 5         | 3 [1.0]  | 3 [1.0]  |  |
| Bile Duct, Cyst                         | 1 [3.0]  |           |          |          |  |
| Hepatocyte, Hypertrophy                 |          | 1 [2.0]   |          |          |  |
| lesentery                               | (10)     | (3)       | (4)      | (2)      |  |
| Fat, Necrosis                           | 9 [2.9]  | 3 [3.0]   | 3 [3.0]  | 1 [3.0]  |  |
| ancreas                                 | (50)     | (50)      | (50)     | (50)     |  |
| Hyperplasia                             | 1 [1.0]  | ` ,       | ` ,      | ,        |  |
| Infiltration Cellular, Mononuclear Cell | 7 [1.0]  |           | 3 [1.0]  | 6 [1.2]  |  |
| Inflammation                            | 2 [2.0]  |           |          |          |  |
| Vacuolization Cytoplasmic               |          |           |          | 1 [2.0]  |  |
| Acinus, Atrophy                         |          | 2 [1.0]   |          | 1 [2.0]  |  |
| Duct, Cyst                              |          |           |          | 1 [4.0]  |  |
| alivary Glands                          | (50)     | (50)      | (50)     | (50)     |  |
| Atrophy                                 |          | 1 [2.0]   |          | 1 [2.0]  |  |
| Infiltration Cellular, Mononuclear Cell | 35 [1.6] | 28 [1.5]  | 39 [1.5] | 40 [1.8] |  |
| Necrosis                                |          | 1 [2.0]   |          |          |  |
| tomach, Forestomach                     | (50)     | (50)      | (50)     | (50)     |  |
| Hyperplasia, Squamous                   | 1 [1.0]  | 1 [2.0]   | 1 [1.0]  | 1 [2.0]  |  |
| Inflammation                            | 2 [1.0]  |           |          |          |  |
| Ulcer                                   |          |           |          | 2 [2.0]  |  |
| tomach, Glandular                       | (50)     | (50)      | (50)     | (50)     |  |
| Cyst                                    |          |           | 1 [1.0]  |          |  |
| Erosion                                 |          | 1 [3.0]   | 1 [1.0]  | 1 [2.0]  |  |
| Inflammation                            | 1 [2.0]  | 1 [1.0]   |          |          |  |
| Mineralization                          |          |           |          | 1 [2.0]  |  |
| Glands, Cyst                            | 1 [1.0]  | 1 [2.0]   |          | 2 [1.5]  |  |
| Glands, Hyperplasia                     | 1 [2.0]  | 1 [3.0]   | 2 [2.5]  |          |  |
| ooth                                    | (31)     | (32)      | (37)     | (35)     |  |
| Dysplasia                               |          |           | 2        | 1        |  |
| Malformation                            | 29 [1.4] | 32 [1.7]  | 36 [1.4] | 33 [1.8] |  |
| Peridontal Tissue, Inflammation         | 7 [1.7]  | 2 [2.0]   | 1 [2.0]  | 9 [2.0]  |  |
| Pulp, Inflammation                      | 2 [1.5]  |           |          |          |  |

#### CARDIOVASCULAR SYSTEM

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Test Type: CHRONIC

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

Date Report Regsted: 08/21/2007

Lab: BAT

Route: SKIN APPLICATION
Species/Strain: MICE/B6C3F1

| B6C3F1 MICE MALE  | 0 MG/KG  | 0.6 MG/KG          | 2 MG/KG  | 6 MG/KG  |  |
|---|----------|--------------------|----------|----------|--|
| Blood Vessel  | (50)     | (50)               | (50)     | (50)     |  |
| Heart   | (50)     | (50)               | (50)     | (50)     |  |
| Cardiomyopathy<br>Infiltration Cellular, Mononuclear Cell | 9 [1.3]  | 6 [1.2]            | 5 [1.0]  | 3 [1.3]  |  |
| Inflammation  | 1 [2.0]  | 1 [1.0]            |          |          |  |
| Mineralization  |          | 1 [1.0]<br>1 [1.0] |          |          |  |
| Capillary, Hyperplasia                                    |          | 7 [1.0]            | 1 [2.0]  |          |  |
| ENDOCRINE SYSTEM  |          |                    |          |          |  |
| Adrenal Cortex  | (50)     | (50)               | (50)     | (50)     |  |
| Cytoplasmic Alteration                                    | , ,      | 1 [2.0]            | ( /      | ,        |  |
| Degeneration  | 1 [4.0]  |                    |          |          |  |
| Hypertrophy   | 22 [1.7] | 16 [1.6]           | 23 [1.5] | 21 [1.8] |  |
| Pigmentation  | 2 [1.5]  |                    |          |          |  |
| Vacuolization Cytoplasmic                                 | 1 [2.0]  | 1 [1.0]            | 1 [1.0]  |          |  |
| Subcapsular, Hyperplasia                                  | 45 [1.8] | 46 [1.8]           | 46 [1.9] | 44 [1.7] |  |
| Zona Fasciculata, Hyperplasia                             | 6 [2.0]  |                    | 1 [1.0]  | 5 [1.4]  |  |
| Adrenal Medulla   | (50)     | (49)               | (50)     | (50)     |  |
| Hyperplasia   |          | 2 [2.5]            |          |          |  |
| Islets, Pancreatic  | (50)     | (50)               | (50)     | (50)     |  |
| Hyperplasia   | 26 [1.5] | 23 [1.4]           | 28 [1.4] | 25 [1.3] |  |
| Parathyroid Gland   | (42)     | (40)               | (43)     | (41)     |  |
| Cyst  |          |                    | 1 [2.0]  |          |  |
| Pituitary Gland   | (50)     | (50)               | (49)     | (50)     |  |
| Cyst  | 3 [1.7]  | 3 [1.7]            | 4 [1.0]  | 6 [1.5]  |  |
| Hyperplasia   | 1 [2.0]  | 1 [2.0]            | 1 [1.0]  |          |  |
| Thyroid Gland   | (50)     | (50)               | (50)     | (50)     |  |
| Cyst  | 1 [2.0]  |                    |          |          |  |
| Infiltration Cellular, Mononuclear Cell                   |          |                    | 1 [1.0]  |          |  |
| C-cell, Hyperplasia                                       |          |                    |          | 1 [2.0]  |  |
| Follicle, Cyst  | 1 [1.0]  |                    | 2 [3.0]  |          |  |
| Follicular Cell, Hyperplasia                              |          | 3 [1.7]            | 2 [2.5]  | 1 [2.0]  |  |

#### **GENERAL BODY SYSTEM**

None

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

Date Report Regsted: 08/21/2007

| B6C3F1 MICE MALE   | 0 MG/KG         | 0.6 MG/KG                  | 2 MG/KG             | 6 MG/KG             |  |
|--|-----------------|----------------------------|---------------------|---------------------|--|
| ENITAL SYSTEM  |                 |                            |                     |                     |  |
| Coagulating Gland  | (0)             | (1)                        | (2)                 | (1)                 |  |
| Hyperplasia  |                 |                            | 1 [2.0]             |                     |  |
| Inflammation   | (50)            | (50)                       | 1 [2.0]             | (50)                |  |
| Epididymis   | (50)            | (50)                       | (50)                | (50)                |  |
| Granuloma Sperm  | 07 [4 4]        | 05 [4 0]                   | 2 [2.5]             | 2 [2.5]             |  |
| Inflammation   | 37 [1.1]        | 35 [1.0]                   | 35 [1.1]            | 34 [1.0]            |  |
| Penis  | (0)             | (2)                        | (0)                 | (0)                 |  |
| Congestion   | (50)            | 1 [2.0]                    | (50)                | (50)                |  |
| Preputial Gland<br>Infiltration Cellular, Mononuclear Cell   | (50)            | (50)                       | (50)                | (50)                |  |
| Inflammation   | 20 [4 4]        | 20 [4 6]                   | 06 [4 4]            | 1 [3.0]             |  |
| Duct, Ectasia  | 38 [1.4]        | 30 [1.6]                   | 26 [1.4]            | 28 [1.4]            |  |
| Prostate   | 2 [2.0]         | 2 [2.5]<br>(50)            | 1 [2.0]<br>(50)     | 4 [2.3]<br>(50)     |  |
| Atrophy  | (50)            | (50)                       | (50)                | 1 [3.0]             |  |
| Infiltration Cellular, Mononuclear Cell                      | 20 [1.2]        | 17 [1.1]                   | 18 [1.1]            | 1 [3.0]<br>14 [1.3] |  |
| Inflammation, Suppurative                                    | 20 [1.2]        | 17 [1.1]                   | 16 [1.1]<br>1 [3.0] | 14 [1.3]            |  |
| Seminal Vesicle  | (50)            | (50)                       | (50)                | (50)                |  |
| Atrophy  | (50)            | (30)                       | (30)                | 1 [3.0]             |  |
| Testes   | (50)            | (50)                       | (50)                | (50)                |  |
| Mineralization   | 1 [2.0]         | (50)                       | 3 [1.0]             | 1 [1.0]             |  |
| Germinal Epithelium, Atrophy                                 | 4 [1.8]         |                            | 3 [2.0]             | 5 [2.2]             |  |
| EMATOPOIETIC SYSTEM  |                 |                            | . []                | - ()                |  |
| Bone Marrow  | (50)            | (50)                       | (50)                | (50)                |  |
| Angiectasis  | 1 [2.0]         |                            |                     |                     |  |
| Myelofibrosis  |                 | 2 [1.5]                    | 2 [1.0]             | 1 [3.0]             |  |
| Myeloid Cell, Hyperplasia                                    | 3 [2.3]         | 4 [2.5]                    | 3 [2.0]             | 3 [2.0]             |  |
| Lymph Node   | (0)             | (1)                        | (3)                 | (1)                 |  |
| Renal, Hyperplasia, Lymphoid                                 |                 |                            | 1 [4.0]             |                     |  |
| Lymph Node, Mandibular                                       | (50)            | (50)                       | (50)                | (50)                |  |
| Atrophy  | 1 [3.0]         |                            |                     | 1 [3.0]             |  |
| Hemorrhage   |                 | 1 [2.0]                    |                     |                     |  |
| Hyperplasia, Lymphoid  | 2 [2.0]         | 1 [2.0]                    |                     | 1 [2.0]             |  |
| r typerplasia, Lymphola                                      |                 |                            |                     |                     |  |
| Hyperplasia, Plasma Cell                                     |                 | 4 [2.0]                    | 1 [2.0]             |                     |  |
| Hyperplasia, Plasma Cell Pigmentation Lymph Node, Mesenteric | 1 [2.0]<br>(50) | 4 [2.0]<br>1 [2.0]<br>(50) | 1 [2.0]<br>(50)     | (50)                |  |

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

## P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

Date Report Regsted: 08/21/2007

| B6C3F1 MICE MALE  | 0 MG/KG  | 0.6 MG/KG                   | 2 MG/KG  | 6 MG/KG                                 |  |
|---|--|-----------------------------|--|---|--|
| Atrophy   | 1 [3.0]  | 1 [2.0]                     | 1 [3.0]  | 1 [2.0]                                 |  |
| Hemorrhage  | 1 [2.0]  | 1                           | 1 [2.0]  | 1 [2.0]                                 |  |
| Hyperplasia, Histiocytic  |  |                             | 1 [2.0]  |   |  |
| Hyperplasia, Lymphoid   | 1 [4.0]  | 2 [3.5]                     | 1 [3.0]  | 1 [2.0]                                 |  |
| Hyperplasia, Plasma Cell  |  | 2 [2.5]                     |  |   |  |
| Inflammation, Suppurative   | 1 [3.0]  |                             |  |   |  |
| Inflammation, Granulomatous   | 1 [3.0]  |                             |  |   |  |
| Spleen  | (50)   | (49)                        | (50)   | (50)                                    |  |
| Hematopoietic Cell Proliferation  | 19 [2.3]   | 13 [2.5]                    | 20 [2.2]   | 13 [2.5]                                |  |
| Hyperplasia, Histiocytic  |  |                             |  | 1 [3.0]                                 |  |
| Hyperplasia, Lymphoid   | 8 [2.1]  | 12 [2.1]                    | 12 [1.8]   | 11 [2.4]                                |  |
| Infiltration Cellular, Polymorphonuclear  |  |                             | 1 [3.0]  |   |  |
| Capsule, Degeneration   | 1 [2.0]  |                             |  |   |  |
| Lymphoid Follicle, Atrophy  | 2 [2.5]  | 4 [2.5]                     |  | 1 [2.0]                                 |  |
| Red Pulp, Atrophy   | 1 [3.0]  | 4 [2.5]                     | 1 [2.0]  | 2 [2.5]                                 |  |
| Thymus  | (47)   | (48)                        | (46)   | (46)                                    |  |
| Atrophy   | 22 [2.5]   | 19 [2.5]                    | 27 [2.4]   | 23 [2.5]                                |  |
| Cyst  |  | 2 [1.0]                     |  | 1 [2.0]                                 |  |
|   |  |                             |  |   |  |
| Skin Hyperplasia Inflammation Inflammation, Chronic Active Ulcer Site Of Application - Dermis, Inflammation, Chronic Active Site Of Application - Epidermis, Hyperplasia Site Of Application - Epidermis, Ulcer | (50)<br>1 [3.0]<br>1 [3.0]<br>4 [1.3]<br>6 [2.0] | (50)<br>1 [1.0]<br>12 [1.0] | (50)<br>1 [2.0]<br>2 [2.0]<br>2 [3.0]<br>7 [1.1]<br>37 [1.1] | (50)<br>10 [1.6]<br>50 [1.6]<br>3 [2.7] |  |
| Skin Hyperplasia Inflammation Inflammation, Chronic Active Ulcer Site Of Application - Dermis, Inflammation, Chronic Active Site Of Application - Epidermis, Hyperplasia  | 1 [3.0]<br>1 [3.0]<br>4 [1.3]                    | 1 [1.0]                     | 1 [2.0]<br>2 [2.0]<br>2 [3.0]<br>7 [1.1]<br>37 [1.1]         | 10 [1.6]<br>50 [1.6]                    |  |
| Skin Hyperplasia Inflammation Inflammation, Chronic Active Ulcer Site Of Application - Dermis, Inflammation, Chronic Active Site Of Application - Epidermis, Hyperplasia Site Of Application - Epidermis, Ulcer | 1 [3.0]<br>1 [3.0]<br>4 [1.3]                    | 1 [1.0]                     | 1 [2.0]<br>2 [2.0]<br>2 [3.0]<br>7 [1.1]<br>37 [1.1]         | 10 [1.6]<br>50 [1.6]                    |  |

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

## P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

Date Report Regsted: 08/21/2007

| B6C3F1 MICE MALE                                      | 0 MG/KG            | 0.6 MG/KG       | 2 MG/KG            | 6 MG/KG         |
|---|--------------------|-----------------|--------------------|-----------------|
| ERVOUS SYSTEM   |                    |                 |                    |                 |
| Brain   | (50)               | (50)            | (50)               | (50)            |
| Hemorrhage<br>Meninges, Infiltration Cellular,        |                    | 1 [2.0]         | 1 [1.0]            | 1 [2.0]         |
| Mononuclear Cell                                      |                    |                 |                    |                 |
| Spinal Cord   | (0)                | (0)             | (0)                | (1)             |
| Atrophy   |                    |                 |                    | 1 [2.0]         |
| ESPIRATORY SYSTEM                                     |                    |                 |                    |                 |
| Lung  | (50)               | (50)            | (50)               | (50)            |
| Hemorrhage<br>Infiltration Cellular, Mononuclear Cell | 1 [3.0]            | 1 [3.0]         |                    | 1 [2.0]         |
| Inflammation  | 6 [1.7]            | 5 [2.2]         |                    | 1 [2.0]         |
| Pigmentation<br>Thrombosis                            | 1 [2.0]<br>1 [2.0] | 1 [1.0]         |                    |                 |
| Alveolar Epithelium, Hyperplasia                      | 6 [1.7]            |                 |                    | 4 [2.8]         |
| Alveolus, Infiltration Cellular, Histiocyte           | 5 [2.2]            | 8 [1.8]         | 2 [2.0]            | 3 [1.3]         |
| Bronchiole, Hyperplasia<br>Nose                       | (50)               | (50)            | 1 [2.0]<br>(50)    | 1 [2.0]<br>(50) |
| Inflammation, Suppurative                             | (30)               | 3 [2.3]         | (30)               | (30)            |
| Polyp, Inflammatory                                   | 1 [3.0]            |                 | 1 [3.0]            |                 |
| Glands, Dilatation<br>Nerve, Atrophy                  |                    |                 | 1 [2.0]<br>1 [2.0] |                 |
| Respiratory Epithelium, Hyperplasia                   |                    |                 | 1 [1.0]            |                 |
| Respiratory Epithelium, Inflammation                  | 7 [1.4]            | 4 [0 0]         | 1 [1.0]            |                 |
| Respiratory Epithelium, Metaplasia<br>Trachea         | (50)               | 1 [3.0]<br>(50) | (50)               | (50)            |
| Inflammation  | ` ,                | (00)            | (00)               | 1 [1.0]         |
| Glands, Cyst  | 1 [1.0]            |                 |                    |                 |
| PECIAL SENSES SYSTEM                                  |                    |                 |                    |                 |
| Eye   | (50)               | (50)            | (50)               | (50)            |
| Atrophy   | ` '                | 1 [4.0]         |                    |                 |
| Cornea, Inflammation<br>Lens, Degeneration            | 1 [3.0]            | 1 [3.0]         |                    | 2 [1.0]         |
| Retina, Atrophy                                       | 2 [4.0]            | 1 [2.0]         |                    |                 |

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

TDMS No. 97003 - 08
Test Type: CHRONIC

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

## P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Date Report Reqsted: 08/21/2007
Time Report Reqsted: 13:25:49

First Dose M/F: 06/25/02 / 06/24/02

| B6C3F1 MICE MALE                        | 0 MG/KG  | 0.6 MG/KG | 2 MG/KG  | 6 MG/KG  |
|---|----------|-----------|----------|----------|
| Harderian Gland                         | (50)     | (50)      | (50)     | (50)     |
| Fibrosis, Focal                         | ()       | 1 [3.0]   | ()       | ()       |
| Infiltration Cellular, Mononuclear Cell | 31 [1.2] | 30 [1.2]  | 26 [1.2] | 30 [1.1] |
| Inflammation                            | 1 [2.0]  |           | 1 [3.0]  |          |
| Epithelium, Hyperplasia                 | 2 [2.0]  | 3 [1.0]   |          | 4 [2.0]  |
| URINARY SYSTEM                          |          |           |          |          |
| Kidney                                  | (50)     | (50)      | (50)     | (50)     |
| Cyst                                    | 3 [1.0]  | 8 [1.8]   | 2 [1.5]  | 6 [1.3]  |
| Hemorrhage                              |          |           |          | 1 [3.0]  |
| Infarct                                 | 4 [1.8]  | 1 [1.0]   | 4 [1.8]  | 3 [1.7]  |
| Infiltration Cellular, Mononuclear Cell | 1 [2.0]  | 1 [3.0]   |          |          |
| Inflammation                            | 2 [3.0]  |           |          |          |
| Mineralization                          | 44 [1.0] | 45 [1.0]  | 46 [1.0] | 43 [1.0] |
| Nephropathy                             | 48 [1.5] | 41 [1.2]  | 48 [1.1] | 48 [1.6] |
| Papilla, Renal Tubule, Necrosis         |          | 1 [3.0]   |          |          |
| Renal Tubule, Hyperplasia               | 12 [1.0] | 12 [1.0]  | 12 [1.0] | 7 [1.0]  |
| Urinary Bladder                         | (50)     | (50)      | (50)     | (50)     |
| Infiltration Cellular, Mononuclear Cell | 22 [1.2] | 19 [1.2]  | 23 [1.0] | 27 [1.2] |
| Transitional Epithelium, Hyperplasia    |          |           | 2 [2.0]  |          |

<sup>\*\*\*</sup> END OF MALE \*\*\*

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH **AVERAGE SEVERITY GRADES[b]**

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Date Report Regsted: 08/21/2007 Time Report Reqsted: 13:25:49

First Dose M/F: 06/25/02 / 06/24/02

Lab: BAT

| B6C3F1 MICE FEMALE                             | 0 MG/KG        | 0.6 MG/KG      | 2 MG/KG       | 6 MG/KG       |  |
|--|----------------|----------------|---------------|---------------|--|
| Disposition Summary                            |                |                |               |               |  |
| Animals Initially in Study                     | 50             | 50             | 50            | 50            |  |
| Early Deaths                                   |                |                |               |               |  |
| Moribund Sacrifice                             | 10             | 11             | 14            | 11            |  |
| Natural Death                                  | 7              | 3              | 6             | 4             |  |
| Survivors                                      |                |                |               |               |  |
| Natural Death                                  |                | 1              |               |               |  |
| Terminal Sacrifice                             | 33             | 35             | 30            | 35            |  |
| Animals Examined Microscopically               | 50             | 50             | 50            | 50            |  |
|  |                |                |               |               |  |
| ALIMENTARY SYSTEM                              |                |                |               |               |  |
| Esophagus                                      | (50)           | (50)           | (50)          | (50)          |  |
| Gallbladder                                    | (50)           | (49)           | (50)          | (50)          |  |
| Cyst   | (33)           | 1 [1.0]        | (00)          | (00)          |  |
| Inflammation                                   |                | 3 [1.7]        | 5 [1.0]       | 2 [1.0]       |  |
| Intestine Small, Duodenum                      | (50)           | (50)           | (50)          | (50)          |  |
| Intestine Small, Jejunum                       | (50)           | (50)           | (50)          | (50)          |  |
| Hyperplasia, Lymphoid                          | , ,            | ,              | 1 [3.0]       | ,             |  |
| Inflammation                                   | 3 [2.7]        | 1 [1.0]        |               |               |  |
| Necrosis, Fatty                                | 1 [3.0]        |                |               |               |  |
| Liver  | (50)           | (50)           | (50)          | (50)          |  |
| Angiectasis                                    |                | 1 [2.0]        | 1 [1.0]       |               |  |
| Basophilic Focus                               | 1              | 3              |               |               |  |
| Clear Cell Focus                               | 14             | 13             | 3             | 8             |  |
| Cyst   | _              | 1 [2.0]        |               | _             |  |
| Eosinophilic Focus                             | 6              | 3              | 4             | 4             |  |
| Fatty Change, Focal                            | 1 [2.0]        |                | 1 [2.0]       | 4.50.01       |  |
| Fatty Change, Diffuse                          | 1 [1.0]        | 4 [0 0]        | 4 [1.5]       | 4 [2.8]       |  |
| Fibrosis                                       | C [4 E]        | 1 [3.0]        | 0.14.01       | 5 [4 O]       |  |
| Hematopoietic Cell Proliferation               | 6 [1.5]        | 3 [1.0]        | 3 [1.0]       | 5 [1.0]       |  |
| Infiltration Cellular, Lymphocyte Inflammation | 25 [4 2]       | 10 11 CN       | 1 [2.0]       | 41 [1 4]      |  |
| Mixed Cell Focus                               | 35 [1.2]<br>13 | 42 [1.2]<br>10 | 37 [1.1]<br>5 | 41 [1.1]<br>7 |  |
| Manager Cell Fucus                             | 10             | 0.10           | 5             | /             |  |

3 [2.0]

4 [1.3]

5

Tension Lipidosis

Necrosis

Pigmentation

8 [1.9]

1 [1.0]

4

4 [2.0]

1 [1.0]

6

2 [2.0]

4 [1.5]

2

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Test Type: CHRONIC

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

Date Report Regsted: 08/21/2007

Lab: BAT

| <del>-</del> -              |
|-----------------------------|
| Route: SKIN APPLICATION     |
| Species/Strain: MICE/B6C3F1 |
|                             |

| B6C3F1 MICE FEMALE                                      | 0 MG/KG  | 0.6 MG/KG           | 2 MG/KG            | 6 MG/KG            |  |
|---|----------|---------------------|--------------------|--------------------|--|
| Vacuolization Cytoplasmic                               |          |                     | 1 [1.0]            |                    |  |
| Hepatocyte, Hypertrophy                                 | 1 [3.0]  |                     |                    |                    |  |
| Mesentery   | (10)     | (9)                 | (6)                | (10)               |  |
| Fat, Necrosis   | 9 [3.0]  | 9 [3.0]             | 6 [3.0]            | 9 [3.0]            |  |
| Pancreas  | (50)     | (50)                | (50)               | (50)               |  |
| Cyst  | 1 [2.0]  | 2 [3.0]             | 47 [4 4]           | 0.4.[4.0]          |  |
| Infiltration Cellular, Mononuclear Cell<br>Inflammation | 17 [1.0] | 15 [1.1]<br>3 [1.7] | 17 [1.1]           | 24 [1.3]           |  |
| Acinus, Atrophy   | 1 [2.0]  | 3 [3.3]             |                    |                    |  |
| Salivary Glands   | (50)     | (50)                | (50)               | (50)               |  |
| Infiltration Cellular, Mononuclear Cell                 | 31 [1.5] | 32 [1.3]            | 33 [1.5]           | 36 [1.5]           |  |
| Stomach, Forestomach                                    | (50)     | (50)                | (50)               | (50)               |  |
| Cyst  | 1 [1.0]  |                     |                    |                    |  |
| Hyperkeratosis  | 1 [2.0]  |                     |                    | 2 [2.5]            |  |
| Hyperplasia, Squamous                                   | 2 [1.5]  |                     |                    | 2 [2.0]            |  |
| Inflammation  |          |                     |                    | 2 [2.0]            |  |
| Ulcer   | (50)     | (50)                | (50)               | 2 [2.0]            |  |
| Stomach, Glandular                                      | (50)     | (50)                | (50)               | (50)               |  |
| Cyst<br>Erosion   | 1 [1.0]  | 2 [2 0]             |                    | 1 [2.0]            |  |
| Mineralization  |          | 2 [2.0]             | 1 [1.0]            | 1 [2.0]<br>1 [2.0] |  |
| Glands, Cyst  | 2 [1.5]  | 5 [1.0]             | 4 [1.0]            | 5 [1.0]            |  |
| Glands, Gyst<br>Glands, Hyperplasia                     | 2 [1.0]  | 1 [2.0]             | <del>-</del> [1.0] | 3 [1.0]            |  |
| Tooth   | (3)      | (4)                 | (4)                | (1)                |  |
| Malformation  | 3 [1.3]  | 3 [1.0]             | 3 [1.0]            | 1 [1.0]            |  |
| Peridontal Tissue, Inflammation                         | 0 [0]    | 1 [2.0]             | 1 [2.0]            | . [0]              |  |
| ARDIOVASCULAR SYSTEM                                    |          |                     |                    |                    |  |
| Blood Vessel  | (50)     | (50)                | (50)               | (50)               |  |
| Hyperplasia   |          | 1 [2.0]             |                    |                    |  |
| Heart   | (50)     | (50)                | (50)               | (50)               |  |
| Cardiomyopathy  | 10 [1.0] | 2 [1.0]             | 9 [1.0]            | 14 [1.0]           |  |
| Infiltration Cellular, Mononuclear Cell                 | 1 [1.0]  |                     | 1 [1.0]            | 1 [2.0]            |  |
| Inflammation  | 4 [0 0]  |                     |                    | 2 [2.0]            |  |
| Mineralization Thrombosis                               | 1 [2.0]  |                     |                    | 1 [2 0]            |  |
| Thrombosis  |          |                     |                    | 1 [3.0]            |  |

#### **ENDOCRINE SYSTEM**

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Test Type: CHRONIC

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

Duct, Cyst

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Date Report Reqsted: 08/21/2007

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

Lab: BAT

| B6C3F1 MICE FEMALE                      | 0 MG/KG  | 0.6 MG/KG | 2 MG/KG  | 6 MG/KG  |  |
|---|----------|-----------|----------|----------|--|
| Adrenal Cortex                          | (50)     | (50)      | (50)     | (50)     |  |
| Hematopoietic Cell Proliferation        | 2 [2.0]  | ()        | ()       | 1 [2.0]  |  |
| Hypertrophy                             | 2 [2.0]  | 2 [1.5]   |          | 1 [2.0]  |  |
| Inflammation                            | 1 [2.0]  |           |          | 1 [3.0]  |  |
| Vacuolization Cytoplasmic               |          |           | 1 [3.0]  | 1 [3.0]  |  |
| Capsule, Necrosis, Fatty                | 1 [2.0]  |           | 1        | []       |  |
| Subcapsular, Hyperplasia                | 50 [2.5] | 49 [2.3]  | 50 [2.2] | 49 [2.3] |  |
| Zona Fasciculata, Hyperplasia           | 2 [1.5]  | 2 [1.5]   | 1 [2.0]  | 1 [2.0]  |  |
| Adrenal Medulla                         | (50)     | (50)      | (50)     | (50)     |  |
| Hyperplasia                             | 1 [2.0]  | 2 [1.0]   | ()       | 3 [2.0]  |  |
| Vacuolization Cytoplasmic               | . [=,]   | -[]       |          | 1 [3.0]  |  |
| Islets, Pancreatic                      | (50)     | (50)      | (50)     | (50)     |  |
| Hyperplasia                             | 4 [1.3]  | ()        | ()       | 1 [1.0]  |  |
| Parathyroid Gland                       | (38)     | (40)      | (40)     | (41)     |  |
| Cyst                                    | (/       | (15)      | (12)     | 1 [1.0]  |  |
| Inflammation, Chronic Active            |          |           |          | 1 [2.0]  |  |
| Pituitary Gland                         | (50)     | (50)      | (50)     | (50)     |  |
| Angiectasis                             | 1 [2.0]  | 2 [2.0]   | 1 [2.0]  | 1 [3.0]  |  |
| Cyst                                    | 1 [1.0]  | 1 [1.0]   | . []     | 2 [1.0]  |  |
| Pars Distalis, Angiectasis              | . []     | 1 [2.0]   |          | _ []     |  |
| Pars Distalis, Hyperplasia              | 9 [1.6]  | 13 [2.2]  | 16 [1.9] | 12 [2.3] |  |
| Pars Intermedia, Hyperplasia            | - []     | 1 [1.0]   | []       | []       |  |
| Thyroid Gland                           | (50)     | (50)      | (50)     | (50)     |  |
| Infiltration Cellular, Mononuclear Cell | 3 [1.0]  | 2 [1.0]   | 3 [1.0]  | 3 [1.3]  |  |
| Inflammation, Suppurative               | 1 [1.0]  | _[]       | 0 [0]    | 0 [0]    |  |
| C-cell, Hyperplasia                     | . []     | 1 [1.0]   |          |          |  |
| Follicle, Cyst                          | 2 [1.5]  | . []      |          |          |  |
| Follicular Cell, Hyperplasia            | 1 [1.0]  | 3 [1.7]   | 1 [2.0]  | 1 [4.0]  |  |
| Follicular Cell, Hypertrophy            | 1 [1.0]  | 9 []      | . [=.0]  | . []     |  |
| ENERAL BODY SYSTEM                      |          |           |          |          |  |
| None                                    |          |           |          |          |  |
| ENITAL SYSTEM                           |          |           |          |          |  |
| Clitoral Gland                          | (50)     | (49)      | (50)     | (50)     |  |
| Inflammation                            | 18 [1.0] | 5 [1.0]   | 2 [1.0]  | 7 [1.1]  |  |

a - Number of animals examined microscopically at site and number of animals with lesion

1 [1.0]

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49

Date Report Regsted: 08/21/2007

First Dose M/F: 06/25/02 / 06/24/02

| B6C3F1 MICE FEMALE   | 0 MG/KG         | 0.6 MG/KG | 2 MG/KG  | 6 MG/KG            |  |
|--|-----------------|-----------|----------|--------------------|--|
| Ovary  | (50)            | (50)      | (50)     | (50)               |  |
| Angiectasis  | ,               | 1 [3.0]   | 3 [2.7]  | 1 [3.0]            |  |
| Cyst   | 7 [2.6]         | 10 [2.2]  | 19 [1.9] | 13 [2.2]           |  |
| Hemorrhage   |                 |           | 1 [2.0]  |                    |  |
| Inflammation, Granulomatous                                    |                 | 1 [2.0]   |          |                    |  |
| Mineralization   |                 | 1 [3.0]   |          |                    |  |
| Pigmentation   |                 | 1 [2.0]   |          |                    |  |
| Thrombosis   | 1 [3.0]         | 1 [3.0]   | 2 [3.0]  |                    |  |
| Periovarian Tissue, Necrosis                                   | 1 [3.0]         |           |          |                    |  |
| Uterus   | (50)            | (50)      | (50)     | (50)               |  |
| Angiectasis  | 2 [2.0]         | 1 [3.0]   |          |                    |  |
| Inflammation, Suppurative                                      |                 |           |          | 1 [4.0]            |  |
| Thrombosis   |                 |           | 1 [2.0]  |                    |  |
| Endometrium, Hyperplasia, Cystic                               | 50 [3.1]        | 49 [2.7]  | 48 [2.7] | 48 [2.9]           |  |
| Vagina   | (0)             | (0)       | (3)      | (0)                |  |
| HEMATOPOIETIC SYSTEM   |                 |           |          |                    |  |
| Bone Marrow  | (50)            | (50)      | (50)     | (50)               |  |
| Angiectasis  |                 | 1 [1.0]   | 1 [3.0]  |                    |  |
| Atrophy, Focal   | 3 [2.0]         | 00.14.43  | 1 [2.0]  | 2 [2.0]            |  |
| Myelofibrosis  | 27 [1.3]        | 29 [1.4]  | 27 [1.3] | 27 [1.4]           |  |
| Myeloid Cell, Hyperplasia                                      | 2 [3.0]         | (0)       | (0)      | 4 [2.0]            |  |
| Lymph Node   | (8)             | (6)       | (6)      | (4)                |  |
| Hyperplasia, Plasma Cell                                       |                 |           | 4 [0 0]  | 1 [3.0]            |  |
| Iliac, Ectasia   |                 | 4 [2 0]   | 1 [2.0]  |                    |  |
| Inguinal, Ectasia  |                 | 1 [2.0]   |          | 4 [0 0]            |  |
| Lumbar, Congestion<br>Lumbar, Ectasia                          |                 | 2 [2 0]   | 2 [3.0]  | 1 [2.0]            |  |
| Lumbar, Hyperplasia, Histiocytic                               |                 | 2 [3.0]   | 2 [3.0]  | 1 [2.0]            |  |
| Mediastinal, Congestion  |                 |           |          | 1 [2.0]<br>1 [3.0] |  |
| Mediastinal, Congestion  Mediastinal, Hyperplasia, Histiocytic |                 |           |          | 1 [2.0]            |  |
| Renal, Ectasia   | 2 [3.0]         |           | 1 [3.0]  | 1 [2.0]            |  |
| Lymph Node, Mandibular   | 2 [3.0]<br>(50) | (50)      | (50)     | (50)               |  |
| Atrophy  | (30)            | (30)      | 1 [3.0]  | (30)               |  |
| Ectasia  |                 | 1 [2.0]   | 1 [3.0]  |                    |  |
| Hyperplasia, Lymphoid  | 1 [3.0]         | 4 [2.3]   |          | 5 [2.4]            |  |
| Hyperplasia, Plasma Cell                                       | 1 [2.0]         | ⊤ اٍک.∪ا  |          | رح. <del>-</del> ا |  |
| Lymph Node, Mesenteric   | (50)            | (50)      | (50)     | (50)               |  |
| Lymph Node, Westing  | (30)            | (30)      | (30)     | (55)               |  |

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

## P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49

Date Report Regsted: 08/21/2007

First Dose M/F: 06/25/02 / 06/24/02

| B6C3F1 MICE FEMALE                              | 0 MG/KG  | 0.6 MG/KG | 2 MG/KG  | 6 MG/KG  |  |
|---|----------|-----------|----------|----------|--|
| Congestion, Chronic                             |          |           |          | 1 [3.0]  |  |
| Hyperplasia, Histiocytic                        |          |           |          | 1 [2.0]  |  |
| Hyperplasia, Lymphoid                           |          | 3 [2.0]   | 3 [2.7]  | 2 [2.0]  |  |
| Hyperplasia, Plasma Cell                        | 1 [2.0]  | 1 [2.0]   | 1 [2.0]  | 2 [2.0]  |  |
| Inflammation                                    |          | 1 [3.0]   |          |          |  |
| Spleen  | (50)     | (50)      | (50)     | (50)     |  |
| Fibrosis  |          | 1 [3.0]   |          |          |  |
| Hematopoietic Cell Proliferation                | 16 [2.4] | 19 [2.2]  | 18 [2.3] | 22 [2.5] |  |
| Hyperplasia, Lymphoid                           | 13 [2.4] | 19 [2.1]  | 19 [2.3] | 16 [2.3] |  |
| Pigmentation                                    | 7 [1.0]  | 5 [1.2]   | 8 [1.3]  | 6 [1.2]  |  |
| Lymphoid Follicle, Atrophy                      | 1 [2.0]  |           | 4 [2.5]  | 1 [2.0]  |  |
| Red Pulp, Atrophy                               | 1 [2.0]  | (47)      | 4 [2.3]  | 3 [2.0]  |  |
| Thymus  | (47)     | (47)      | (48)     | (50)     |  |
| Atrophy   | 11 [2.5] | 11 [2.4]  | 13 [2.8] | 17 [2.8] |  |
| Hyperplasia, Atypical                           |          | 1 [2.0]   | 7 [0 4]  | 4 [2 2]  |  |
| Hyperplasia, Lymphoid                           |          | 5 [2.0]   | 7 [2.1]  | 4 [2.3]  |  |
| INTEGUMENTARY SYSTEM                            |          |           |          |          |  |
| Mammary Gland                                   | (50)     | (49)      | (50)     | (50)     |  |
| Hyperplasia                                     | (==)     | (==)      | 1 [1.0]  | ()       |  |
| Skin  | (50)     | (50)      | (50)     | (50)     |  |
| Site Of Application - Dermis,                   |          | 9 [1.4]   | 30 [1.3] | 28 [1.4] |  |
| Inflammation, Chronic Active                    |          | 40 [4 5]  | 27 [4 2] | 40 [4 5] |  |
| Site Of Application - Epidermis,<br>Hyperplasia |          | 12 [1.5]  | 37 [1.2] | 49 [1.5] |  |
| Site Of Application - Epidermis, Ulcer          |          | 1 [3.0]   | 2 [2.5]  | 3 [3.0]  |  |
| Subcutaneous Tissue, Inflammation,              |          | 1 [3.0]   | رد.ی     | 1 [3.0]  |  |
| Granulomatous                                   |          |           |          | 1 [3.0]  |  |
| Subcutaneous Tissue, Necrosis, Fatty            |          |           |          | 1 [3.0]  |  |
| Subcutaneous Tissue, Pigmentation               |          |           |          | 1 [1.0]  |  |
|   |          |           |          | . [•]    |  |
| MUSCULOSKELETAL SYSTEM                          |          |           |          |          |  |
| Bone  | (50)     | (50)      | (50)     | (50)     |  |
| Arthrosis                                       | (00)     | 1 [3.0]   | (00)     | (55)     |  |
| Osteopetrosis                                   |          | . []      |          | 1 [3.0]  |  |
| Skeletal Muscle                                 | (1)      | (1)       | (1)      | (1)      |  |
|   | ` '      | ` '       | ` '      | ` '      |  |

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

## P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49 First Dose M/F: 06/25/02 / 06/24/02

Date Report Regsted: 08/21/2007

| B6C3F1 MICE FEMALE  | 0 MG/KG             | 0.6 MG/KG           | 2 MG/KG             | 6 MG/KG             |  |
|---|---------------------|---------------------|---------------------|---------------------|--|
| NERVOUS SYSTEM  |                     |                     |                     |                     |  |
| Brain   | (50)                | (50)                | (50)                | (50)                |  |
| Hemorrhage<br>Pigmentation  |                     |                     | 1 [2.0]             | 1 [1.0]             |  |
| Neuron, Necrosis  |                     |                     | . []                | 1 [1.0]             |  |
| RESPIRATORY SYSTEM  |                     |                     |                     |                     |  |
| Lung  | (50)                | (50)                | (50)                | (50)                |  |
| Hemorrhage<br>Inflammation  | 1 [3.0]             | 2 [3.0]<br>1 [1.0]  |                     | 2 [3.0]<br>3 [2.0]  |  |
| Pigmentation  |                     |                     |                     | 1 [1.0]             |  |
| Alveolar Epithelium, Hyperplasia<br>Alveolus, Infiltration Cellular, Histiocyte | 1 [2.0]<br>2 [2.5]  | 1 [2.0]<br>1 [2.0]  | 2 [2.0]             | 2 [2.0]<br>1 [1.0]  |  |
| Mediastinum, Inflammation   | رد.ن                | 1 [3.0]             |                     | 1 [1.0]             |  |
| Serosa, Fibrosis<br>Nose  | (50)                | (50)                | 1 [3.0]             | (50)                |  |
| Glands, Cyst  | (50)                | (50)                | (50)                | (50)<br>1 [1.0]     |  |
| Respiratory Epithelium, Inflammation  | 2 [2.0]             | (50)                | (50)                |                     |  |
| Trachea   | (50)                | (50)                | (50)                | (50)                |  |
| SPECIAL SENSES SYSTEM   |                     |                     |                     |                     |  |
| Eye   | (50)                | (50)                | (50)                | (50)                |  |
| Cornea, Inflammation, Suppurative Lens, Degeneration                            | 2 [3.0]             | 1 [3.0]<br>2 [2.5]  |                     |                     |  |
| Optic Nerve, Degeneration   |                     |                     | 1 [2.0]             |                     |  |
| Harderian Gland   | (50)                | (50)                | (50)                | (50)                |  |
| Infiltration Cellular, Mononuclear Cell<br>Epithelium, Hyperplasia              | 31 [1.1]<br>1 [2.0] | 33 [1.1]<br>1 [3.0] | 36 [1.2]<br>4 [1.0] | 36 [1.1]<br>3 [1.7] |  |
|   | 1                   | F1                  | 1                   |                     |  |
| JRINARY SYSTEM  |                     |                     |                     |                     |  |
| Kidney  | (50)                | (50)                | (50)                | (50)                |  |
| Accumulation, Hyaline Droplet Amyloid Deposition                                | 2 [2.5]             | 1 [4.0]             | 1 [3.0]             |                     |  |
| Cyst  | ر د.∪]              |                     | 1 [1.0]             |                     |  |

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

TDMS No. 97003 - 08
Test Type: CHRONIC

**Route: SKIN APPLICATION** 

Species/Strain: MICE/B6C3F1

## P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

1,2-Dibromo-2,4-dicyanobutane CAS Number: 35691-65-7

Time Report Reqsted: 13:25:49
First Dose M/F: 06/25/02 / 06/24/02

Date Report Regsted: 08/21/2007

| B6C3F1 MICE FEMALE                      | 0 MG/KG  | 0.6 MG/KG        | 2 MG/KG  | 6 MG/KG  |  |
|---|----------|------------------|----------|----------|--|
| Degeneration                            |          |                  |          | 1 [2.0]  |  |
| Infarct                                 | 3 [2.3]  | 2 [2.5]          | 8 [1.8]  | 4 [2.0]  |  |
| Inflammation                            | 2 [2.0]  |                  |          |          |  |
| Metaplasia, Osseous                     |          | 1 [2.0]          |          | 1 [2.0]  |  |
| Mineralization                          | 14 [1.2] | 8 [1.0]          | 4 [1.0]  | 3 [1.0]  |  |
| Nephropathy                             | 16 [1.3] | 15 [1.0]         | 16 [1.3] | 18 [1.1] |  |
| Pigmentation                            |          |                  | 1 [1.0]  |          |  |
| Pelvis, Dilatation                      |          |                  | 1 [2.0]  |          |  |
| Renal Tubule, Hyperplasia               | 1 [1.0]  | 2 [1.5]          |          |          |  |
| Renal Tubule, Necrosis                  | 1 [3.0]  | - <del>-</del>   |          |          |  |
| Urinary Bladder                         | (50)     | (50)             | (50)     | (50)     |  |
| Infiltration Cellular, Mononuclear Cell | 33 [1.8] | 37 [1.3 <u>]</u> | 38 [1.5] | 38 [1.4] |  |

<sup>\*\*\*</sup> END OF REPORT \*\*\*

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)